

# STIC Search Report Biotech-Chem Library

### STIC Database Tracking Number: 168608

TO: Paul Martin

Location: 4b71 / 3c18

Tuesday, October 18, 2005

**Art Unit: 1655** 

Phone: 571-272-3348

Serial Number: 10 / 748335

From: Jan Delaval

**Location: Biotech-Chem Library** 

Remsen 1a51

Phone: 571-272-2504

jan.delaval@uspto.gov

# Search Notes





### Scientific and Technical Information Center

## SEARCH REQUEST FORM

Requester's Full Name: Aul	MUTTO E	xaminer # : <u> </u>	10/14/05
Art Unit: 1655 Phone	Number: 2- 3348	Serial Number: 10 7483	35
Location (Bldg/Room#):	(Mailbox #): Res	ults Format Preferred (circle): PA	PER DISK
*********	*********	***********	*****
To ensure an efficient and quality search,	please attach a copy of the cover s	heet, claims, and abstract or fill out the fo	ollowing:
Title of Invention: Thent fich it is your live to be in the Inventors (please provide full names):	ion of Motive site:	alibitule of gly cosyltians	eferciel wing
Inventors (please provide full names):	- surania wilker	Kahne, Paniel Kahne	
Earliest Priority Date: 12 (30)	02		
Search Topic: Please provide a detailed statement of the so elected species or structures, keywords, synd Define any terms that may have a special m	onyms, acronyms, and registry num	bers, and combine with the concept or utili	rched. Include the ty of the invention.
*For Sequence Searches Only* Please incl appropriate serial number.			ers) along with the
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100	Bibliographic	In-house sequence system	s ·
Date Searcher Picked Up: LO   1860 V		Commercial Oligomer	Score/Length
Date Completed: (0)/19/00	Litigation	Interference SPDI Other (specify)	Encode/Transl
Searcher Prep & Review Time:	Fulltext		•
Online Time:	Other		



# STIC SEARCH RESULTS FEEDBACK FORM

### Biotech-Chem Library

Questions about the scope or the results of the search? Contact the searcher or contact:

Mary Hale, Information Branch Supervisor 308-4258, CM1-1E01

Voluntary Results Feedback Form
> I am an examiner in Workgroup: Example: 1610
> Relevant prior art found, search results used as follows:
☐ 102 rejection
☐ 103 rejection
☐ Cited as being of interest.
Helped examiner better understand the invention.
Helped examiner better understand the state of the art in their technology.
Types of relevant prior art found:
☐ Foreign Patent(s)
<ul> <li>Non-Patent Literature</li> <li>(journal articles, conference proceedings, new product announcements etc.)</li> </ul>
> Relevant prior art not found:
Results verified the lack of relevant prior art (helped determine patentability).
Results were not useful in determining patentability or understanding the invention.
Comments:

Drop off or send completed forms to STIC/Blotech-Chem Library CM1 - Circ. Desk



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L36

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(FILE 'HOME' ENTERED AT 15:03:15 ON 18 OCT 2005)
                SET COST OFF
     FILE 'HCAPLUS' ENTERED AT 15:03:41 ON 18 OCT 2005
              1 S US20050142629/PN OR US2003-784335#/AP, PRN
L1
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            117 S E3-E7
L2
                E KAHNE S/AU
              8 S E4,E5
L3
                E WALKER S/AU
            210 S E3
L4
L5
             20 S E17
                E WALKER SUE/AU
             48 S E21
1.6
                SEL RN L1
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L7
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L8
L9
              2 S L8 AND OC5/ES
L10
              1 S L9 AND NCNC3/ES
          75563 S (OC5-C6-C6 OR OC4-OC5-C6-C6-C6)/ES
L12
           1637 S L11 AND OC5/ES
L13
            103 S L12 AND (OC4 AND NCNC3)/ES
             91 S L13 AND P/ELS
L14
L15
             28 S L14 AND 8/NR
L16
             17 S L15 NOT S/ELS
              1 S L16 AND C38H38N4O23P2
L17
              2 S L8 AND OC4-OC5-C6-C6-C6/ES NOT L9
L18
              1 S L18 NOT N/ELS
L19
              1 S 58-98-0
L20
                E N-ACT5EYLGLUCOSAMINE/CN
                E N-ACETYLGLUCOSAMINE/CN
              1 S E3
L21
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L23
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              0 S L19 AND L20 AND L21
L24
L25
           6310 S L19
L26
          25115 S FLUORESCEIN?
          25589 S L25, L26
L27
          2084 S L20
L28
          16687 S UDP OR URID? (L) DIHYDROGEN (L) ? PHOSPH?
L29
          16657 S UDP OR URID? (L) ?PHOSPH? (L) GLUCORON?
L30
          . 4962 S URID? (L) DIPHOSPH?
L31
             40.S L27 AND L28-L31
L32
           5969 S L21
L33
          25484 S ?ACETYLGLUCOSAMIN? OR ?ACETY? (L) ?GLUCOSAMIN?
T.34
L35
              9 S L32 AND L33, L34
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FILE 'HCAPLUS' ENTERED AT 15:25:47 ON 18 OCT 2005

1 S 528-04-1

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1276 S L36
L37
              7 S L37 AND L27
L38
              1 S L38 NOT L35
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                SEL DN AN L35 2 4 8 9
              4 S E1-E12 AND L35
L40
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L41
              5 S L23, L41
L42
              2 S L1-L6 AND L42
L43
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L44
L45
              2 S L44 AND GLCNAC
T.46
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L47
              1 S L17
=> fil uspatall
FILE 'USPATFULL' ENTERED AT 15:29:46 ON 18 OCT 2005
CA INDEXING COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)
FILE 'USPAT2' ENTERED AT 15:29:46 ON 18 OCT 2005
CA INDEXING COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)
=> d 147 bib abs hitstr
L47
    ANSWER 1 OF 1 USPATFULL on STN
AN
       2005:165205 USPATFULL
ΤI
       Identification of active-site inhibitors of glycosyltransferases using a
       generalizable high-throughput screen
IN
       Kahne, Suzanne Walker, Princeton, NJ, UNITED STATES
       Kahne, Daniel, Princeton, NJ, UNITED STATES
PΙ
       US 2005142629
                               20050630
                          A1
AΙ
       US 2003-748335
                               20031230 (10)
                          A1
DT
       Utility
FS
       APPLICATION
       Patrick H. Higgins, Mathews, Collins, Shepherd & McKay, Suite 306, 100
LREP
       Thanet Circle, Princeton, NJ, 08540, US
       Number of Claims: 20
CLMN
       Exemplary Claim: 1
ECL
       9 Drawing Page(s)
LN.CNT 1398
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       A method is described for identifying a compound that modulates the
       ability of a glycosyltransferase to bind a substrate comprising
       combining a glycosyltransferase, a labeled substrate, and a compound, in
       a reaction vessel, under conditions known to be suitable for the
       glycosyltransferase to bind the labeled substrate, measuring an amount
       of labeled substrate bound to the glycosyltransferase, and comparing the
       amount to a standardized amount to identify a relative increase or
       decrease in substrate bound glycosyltransferase, thereby identifying a
       compound that modulates the ability of the glycosyltransferase to bind
       the substrate. A composition comprising an effective amount of a
       compound.of Formula I (the substituents of which are described herein),
       or a stereoisomer, or pharmaceutically acceptable salt thereof, that
       inhibits the ability of a glycosyltransferase to bind a substrate, in a
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### CAS INDEXING IS AVAILABLE FOR THIS PATENT.

pharmaceutically acceptable carrier is provided

### IT 608143-47-1P

(glycosyltransferases inhibitor screening and use for controlling growth of bacteria)

##STR1##

RN 608143-47-1 USPATFULL

CN

Uridine 5'-(trihydrogen diphosphate), P'-[2-deoxy-2-[[[[(3',6'-dihydroxy-3oxospiro[isobenzofuran-1(3H),9'-[9H]xanthen]-5yl)carbonyl]amino]acetyl]amino]-α-D-glucopyranosyl] ester (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

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FILE COVERS 1907 - 18 Oct 2005 VOL 143 ISS 17 FILE LAST UPDATED: 17 Oct 2005 (20051017/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

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L59 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN
    2005:572457 HCAPLUS
AN
DN
    143:90985
    Entered STN: 01 Jul 2005
ED
    Identification of active-site inhibitors of glycosyltransferases using a
TI
    generalizable high-throughput screen
    Kahne, Suzanne Walker; Kahne, Daniel
IN
PΑ
SO
    U.S. Pat. Appl. Publ., 26 pp.
    CODEN: USXXCO
DT
    Patent
LΑ
    English
TC
    ICM C12Q001-48
INCL 435015000
    1-5 (Pharmacology)
    Section cross-reference(s): 7, 33, 63
FAN.CNT 1
    PATENT NO.
                      KIND DATE
                                       APPLICATION NO.
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                      ____
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                      A1
    US 2005142629
                              20050630
                                         US 2003-748335
                                                             20031230 <--
PRAI US 2003-748335
                             20031230
CLASS
PATENT NO.
             CLASS PATENT FAMILY CLASSIFICATION CODES
 US 2005142629 ICM C12Q001-48
               INCL 435015000
US 2005142629 NCL
                     435/015.000
OS MARPAT 143:90985
AB
    A method is described for identifying a compound that modulates the ability
    of a glycosyltransferase to bind a substrate, comprising combining a
    glycosyltransferase, a labeled substrate, and a compound, in a reaction
    vessel, under conditions known to be suitable for the qlycosyltransferase
    to bind the labeled substrate, measuring an amount of labeled substrate
    bound to the glycosyltransferase, and comparing the amount to a standardized
    amount to identify a relative increase or decrease in substrate bound
    glycosyltransferase, thereby identifying a compound that modulates the
    ability of the glycosyltransferase to bind the substrate. A composition
    comprising an effective amount of a compound that inhibits the ability of a
    glycosyltransferase to bind a substrate, in a pharmaceutically acceptable
    carrier, is also provided. The invention further provides methods for
    controlling the growth of bacteria using the compds. of the invention.
    Compds. of the invention include e.g. 5-(4-tert-butylbenzylidene)-3-(4-
    methylpiperidin-1-ylmethyl)-2-thioxothiazolidin-4-one. Preparation of a
    fluoresceinated UDP-N-acetylglucosamine analog
    is included.
    high throughput screen glycosyl transferase inhibitor antibacterial;
    thioxothiazolidinone deriv glycosyl transferase inhibitor antibacterial;
    UDP acetylglucosamine analog prepn glycosyl transferase
    inhibitor screening
IT
    Crystal structure
       (MurG-bound UDP-GlcNAc; glycosyltransferases
       inhibitor screening and use for controlling growth of bacteria)
IT
    Peptidoglycans
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
       (bacterial peptidoglycan synthesis; glycosyltransferases inhibitor
       screening and use for controlling growth of bacteria)
ΙT
    Infection
       (bacterial; glycosyltransferases inhibitor screening and use for
       controlling growth of bacteria)
```

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IT
    Antibacterial agents
    Chromophores
    Drug screening
    Dyes
    Fluorescent substances
        (glycosyltransferases inhibitor screening and use for controlling
       growth of bacteria)
IT
     Enzymes, biological studies
    Radionuclides, biological studies
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (glycosyltransferases inhibitor screening and use for controlling
        growth of bacteria)
     58-98-0, UDP, biological studies 146-91-8, GDP
IT
     491-97-4, TDP 528-04-1 9023-27-2, Gene MurA enzyme
     9033-07-2, Glycosyltransferase 60976-26-3, MurG glycosyltransferase
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (glycosyltransferases inhibitor screening and use for controlling
        growth of bacteria)
     58-64-0, ADP, biological studies 58-97-9, UMP, biological
IT
     studies 2321-07-5, Fluorescein
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (glycosyltransferases inhibitor screening and use for controlling
        growth of bacteria)
IT
     608143-47-1P
     RL: BUU (Biological use, unclassified); SPN (Synthetic preparation); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (qlycosyltransferases inhibitor screening and use for controlling
        growth of bacteria)
IT
                  292168-98-0D, stereoisomers and salts
                                                           292169-21-2
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                                             292170-21-9
                                                           292170-21-9D,
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                          310421-16-0D, stereoisomers and salts
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                          347385-19-7D, stereoisomers and salts
     347387-81-9D, stereoisomers and salts
                                             347389-31-5
                                                          347389-31-5D,
                                             350693-04-8D, stereoisomers and
     stereoisomers and salts
                              350693-04-8
                                                                   745037-23-4
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                         381708-09-4D, stereoisomers and salts
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                                             745037-25-6D, stereoisomers and
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            856570-62-2
                         856570-62-2D, stereoisomers and salts
                                                                   856570-63-3
     856570-63-3D, stereoisomers and salts
                                             856570-64-4
                                                           856570-64-4D,
     stereoisomers and salts
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (glycosyltransferases inhibitor screening and use for controlling
        growth of bacteria)
IT
     19286-16-9
                 108549-23-1 117548-22-8
     RL: RCT (Reactant); RACT (Reactant or reagent)
```

(glycosyltransferases inhibitor screening and use for controlling growth of bacteria)

IT 856570-65-5P 856570-66-6P 856570-67-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(glycosyltransferases inhibitor screening and use for controlling growth of bacteria)

IT 58-98-0, UDP, biological studies 528-04-1

RL: BSU (Biological study, unclassified); BIOL (Biological study) (glycosyltransferases inhibitor screening and use for controlling growth of bacteria)

RN 58-98-0 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate) (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 528-04-1 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate), P'-[2-(acetylamino)-2-deoxy- $\alpha$ -D-glucopyranosyl] ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT 58-97-9, UMP, biological studies 2321-07-5,

Fluorescein

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(glycosyltransferases inhibitor screening and use for controlling growth of bacteria)

RN 58-97-9 HCAPLUS

CN 5'-Uridylic acid (8CI, 9CI) (CA INDEX NAME)

RN 2321-07-5 HCAPLUS

CN Spiro[isobenzofuran-1(3H),9'-[9H]xanthen]-3-one, 3',6'-dihydroxy- (9CI) (CA INDEX NAME)

IT 608143-47-1P

RL: BUU (Biological use, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses) (glycosyltransferases inhibitor screening and use for controlling growth of bacteria)

RN 608143-47-1 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate), P'-[2-deoxy-2-[[[[(3',6'-dihydroxy-3-oxospiro[isobenzofuran-1(3H),9'-[9H]xanthen]-5yl)carbonyl]amino]acetyl]amino]-α-D-glucopyranosyl] ester (9CI) (CA
INDEX NAME)

### IT 117548-22-8

RL: RCT (Reactant); RACT (Reactant or reagent)
 (glycosyltransferases inhibitor screening and use for controlling
 growth of bacteria)

RN 117548-22-8 HCAPLUS

CN 2,5-Pyrrolidinedione, 1-[[[3',6'-dihydroxy-3-oxospiro[isobenzofuran-1(3H),9'-[9H]xanthen]-5(or 6)-yl]carbonyl]oxy]- (9CI) (CA INDEX NAME)

### IT 856570-67-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(glycosyltransferases inhibitor screening and use for controlling growth of bacteria)

RN 856570-67-7 HCAPLUS

CN  $\alpha$ -D-Glucopyranose, 2-deoxy-2-[[[[(3',6'-dihydroxy-3-

oxospiro[isobenzofuran-1(3H),9'-[9H]xanthen]-5yl)carbonyl]amino]acetyl]amino]-, 1-(dihydrogen phosphate) (9CI) (CA
INDEX NAME)

Absolute stereochemistry.

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L59 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN
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AN 2004:354961 HCAPLUS

DN 140:370523

ED Entered STN: 30 Apr 2004

TI Synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into the protein

IN Schultz, Peter G.; Wang, Lei; Zhang, Zhiwen

PA The Scripps Research Institute, USA

SO PCT Int. Appl., 103 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM C07K

CC 6-3 (General Biochemistry)

FAN.CNT 2

FAN. CNT 2																			
PAT	TENT NO.			KIND DATE		APPLICATION NO.						DATE							
WO				A2 20040429			0429	WO 2003-US32870						20031015					
												20031013							
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH.	CN.		
		LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NI,	NO,	NZ,	OM,		
															ТJ,	TM,	TN,		
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													20031015						
EP																			
	R:																PΤ,		
							RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	SK			
US	2005181471				A1		2005	0818	US 2005-94677						20050329				
US	2005	1866!	56		A1	A1 20050825				US 2005-93798						20050329			
US	2005		A1		2005	0825	US 2005-94676						20050329						
	CA US US EP	PATENT WO 2004 WO 2004 W:  RW:  CA 2500 US 2004 US 6927 EP 1558 R:  US 2005 US 2005	PATENT NO.  WO 20040356  W: AE,  CO,  GM,  LS,  PG,  TR,  RW: GH,  KG,  FI,  S0041381  US 6927042  EP 1558747  R: AT,  IE,  US 20051814  US 20051866	PATENT NO.  WO 2004035605  W: AE, AG, CO, CR, GM, HR, LS, LT, PG, PH, TR, TT, RW: GH, GM, KG, KZ, FI, FR, BF, BJ, CA 2500653. US 2004138106 US 6927042 EP 1558747 R: AT, BE, IE, SI, US 2005181471 US 2005186656	PATENT NO.  WO 2004035605  W: AE, AG, AL, CO, CR, CU, GM, HR, HU, LS, LT, LU, PG, PH, PL, TR, TT, TZ, RW: GH, GM, KE, KG, KZ, MD, FI, FR, GB, BF, BJ, CF, CA 2500653. US 2004138106 US 6927042 EP 1558747 R: AT, BE, CH, IE, SI, LT, US 2005181471 US 2005186656	PATENT NO. KINI  WO 2004035605 A2  WO 2004035605 A3  W: AE, AG, AL, AM, CO, CR, CU, CZ, GM, HR, HU, ID, LS, LT, LU, LV, PG, PH, PL, PT, TR, TT, TZ, UA, RW: GH, GM, KE, LS, KG, KZ, MD, RU, FI, FR, GB, GR, BF, BJ, CF, CG, CA 2500653 AA  US 2004138106 A1 US 6927042 B2 EP 1558747 A2 R: AT, BE, CH, DE, IE, SI, LT, LV, US 2005181471 A1 US 2005186656 A1	PATENT NO. KIND  WO 2004035605 A2  WO 2004035605 A3  W: AE, AG, AL, AM, AT, CO, CR, CU, CZ, DE, GM, HR, HU, ID, IL, LS, LT, LU, LV, MA, PG, PH, PL, PT, RO, TR, TT, TZ, UA, UG, RW: GH, GM, KE, LS, MW, KG, KZ, MD, RU, TJ, FI, FR, GB, GR, HU, BF, BJ, CF, CG, CI, CA 2500653 AA  US 2004138106 A1 US 6927042 B2 EP 1558747 A2  R: AT, BE, CH, DE, DK, IE, SI, LT, LV, FI, US 2005181471 A1 US 2005186656 A1	PATENT NO. KIND DATE  WO 2004035605 A2 2004  WO 2004035605 A3 2005  W: AE, AG, AL, AM, AT, AU,  CO, CR, CU, CZ, DE, DK,  GM, HR, HU, ID, IL, IN,  LS, LT, LU, LV, MA, MD,  PG, PH, PL, PT, RO, RU,  TR, TT, TZ, UA, UG, US,  RW: GH, GM, KE, LS, MW, MZ,  KG, KZ, MD, RU, TJ, TM,  FI, FR, GB, GR, HU, IE,  BF, BJ, CF, CG, CI, CM,  CA 2500653 AA 2004  US 2004138106 A1 2004  US 6927042 B2 2005  R: AT, BE, CH, DE, DK, ES,  IE, SI, LT, LV, FI, RO,  US 2005181471 A1 2005  US 2005186656 A1 2005	PATENT NO. KIND DATE  WO 2004035605 A2 20040429  WO 2004035605 A3 20050512  W: AE, AG, AL, AM, AT, AU, AZ,  CO, CR, CU, CZ, DE, DK, DM,  GM, HR, HU, ID, IL, IN, IS,  LS, LT, LU, LV, MA, MD, MG,  PG, PH, PL, PT, RO, RU, SC,  TR, TT, TZ, UA, UG, US, UZ,  RW: GH, GM, KE, LS, MW, MZ, SD,  KG, KZ, MD, RU, TJ, TM, AT,  FI, FR, GB, GR, HU, IE, IT,  BF, BJ, CF, CG, CI, CM, GA,  CA 2500653 AA 20040429  US 2004138106 A1 20040715  US 6927042 B2 20050809  EP 1558747 A2 20050803  R: AT, BE, CH, DE, DK, ES, FR,  IE, SI, LT, LV, FI, RO, MK,  US 2005181471 A1 20050818  US 2005186656 A1 20050825	PATENT NO. KIND DATE  WO 2004035605 A2 20040429 WO 2004035605 A3 20050512  W: AE, AG, AL, AM, AT, AU, AZ, BA,	PATENT NO. KIND DATE APPL  WO 2004035605 A2 20040429 WO 2  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, GM, HR, HU, ID, IL, IN, IS, JP, KE, LS, LT, LU, LV, MA, MD, MG, MK, MN, PG, PH, PL, PT, RO, RU, SC, SD, SE, TR, TT, TZ, UA, UG, US, UZ, VC, VN, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, FI, FR, GB, GR, HU, IE, IT, LU, MC, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, CA 2500653 AA 20040429 CA 2: US 2004138106 A1 20040715 US 2: US 6927042 B2 20050809 EP 1558747 A2 20050803 EP 2: CA 2505181471 A1 20050818 US 2: US 2005181471 A1 20050818 US 2: US 2005186656 A1 20050825 US 2:  ON 2005181471 A1 2005082	PATENT NO. KIND DATE APPLICATE  WO 2004035605 A2 20040429 WO 2003-1  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, CA 2500653.  US 2004138106 A1 20040715 US 2003-6  US 6927042 B2 20050809  EP 1558747 A2 20050803 EP 2003-7  R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, US 2005181471 A1 20050818 US 2005-5	PATENT NO. KIND DATE APPLICATION 1  WO 2004035605 A2 20040429 WO 2003-US32  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, CA 2500653. AA 20040429 CA 2003-2500 US 2004138106 A1 20040715 US 2003-6869 US 6927042 B2 20050809 EP 1558747 A2 20050803 EP 2003-7776 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, US 2005181471 A1 20050818 US 2005-94675 US 2005186656 A1 20050825 US 2005-93795	PATENT NO. KIND DATE APPLICATION NO.  WO 2004035605 A2 20040429 WO 2003-US32870  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, CA 2500653.  US 2004138106 A1 20040715 US 2003-686944  US 6927042 B2 20050809  EP 1558747 A2 20050803 EP 2003-777634  R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, US 2005181471 A1 20050818 US 2005-94677  US 2005186656 A1 20050825 US 2005-93798	PATENT NO. KIND DATE APPLICATION NO.  WO 2004035605 A2 20040429 WO 2003-US32870  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, CA 2500653 US 2004138106 A1 20040715 US 2003-686944  US 6927042 B2 20050809  EP 1558747 A2 20050803 EP 2003-777634  R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, US 2005181471 A1 20050818 US 2005-93798	PATENT NO. KIND DATE APPLICATION NO. D.  WO 2004035605 A2 20040429 WO 2003-US32870 2  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, CA 2500653 A2 20050809  EP 1558747 A2 20050803 EP 2003-777634 200508181471 A1 20050818 US 2005-94677 200508181471 A1 20050818 US 2005-94677 200508181471 A1 20050818 US 2005-93798 20050818 CS 2005186656 A1 20050825 US 2005-93798 20050809	PATENT NO. KIND DATE APPLICATION NO. DATE  WO 2004035605 A2 20040429 WO 2003-US32870 20031 WO 2004035605 A3 20050512  W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, CA 2500653 A2 20050809  EP 1558747 A2 20050803 EP 2003-777634 200310 S 6927042 B2 20050809 EP 1558747 A2 20050803 EP 2003-777634 200310 S 6927042 B2 20050809 EP 1558747 A2 20050803 EP 2003-777634 200310 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK US 2005181471 A1 20050818 US 2005-94677 2005050050000000000000000000000000000		

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                       514/008.000
EP 1558747 ECLA C12P021/00B
US 2005181471 NCL 435/068.100; 530/395.000; 530/409.000
US 2005186656 NCL
                      435/068.100; 435/193.000; 530/395.000
              NCL
US 2005186657
                       435/068.100; 435/193.000; 530/395.000
US 2005209133
               NCL
                       514/008.000; 530/322.000
    Methods for glycosidating proteins to give novel positions and patterns of
AΒ
    glycosidation are described. One method involves incorporating an
    unnatural amino acid containing a reactive group into a protein and attaching
     one or more saccharide moieties to the unnatural amino acid. Another
    method involves incorporating an unnatural amino acid that includes a
     saccharide moiety into a protein. Proteins made by both methods can be
     further modified with addnl. sugars. Methods of introducing ketoamino
     acids into proteins during protein synthesis by means of tRNA variants
     charged with the amino acid and aminoacyl-tRNA synthetase derivs. capable
    of charging the tRNAs with ketoaminoacids are described. The tRNA
    recognizes a codon such as a stop codon, a rare codon, or a
    tetranucleotide or longer sequence that is rare in the gene of interest.
    A mutant Methanococcus jannaschii tyrosyl tRNA synthetase that could
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and biotin hydrazide at the corresponding sites.

ST protein glycosidation translation amino acid analog reactive group; ketoamino acid protein synthesis tRNA synthetase variant; amino acid analog protein synthesis tRNA synthetase variant

tRNA. The protein could be modified with fluorescein hydrazide

IT tRNA

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

suppress amber mutations in a chloramphenicol acetyltransferase gene was selected and screened for growth on chloramphenicol in the presence

p-acetyl-L-phenylalanine. Translation of genes containing amber mutations in the presence of this synthetase resulted in the introduction of the keto amino acid at the specific sites in the presence of an amer suppressor

(amber suppressor, incorporation of amino acid analogs using; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Codons

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(amber, suppression of, incorporation of amino acid analogs using;
synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Nucleophiles

(amino acid analogs as, for glycosidation; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Glycosides

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(amino acid, incorporation into proteins of; synthetic glycosylation of

proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Amino acids, biological studies

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(analogs, incorporation into proteins of; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Escherichia coli

(expression host; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT DNA sequences

(for tyrosyl tRNA synthetase variants of Methanococcus jannaschii; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Amino acids, biological studies

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(glycosides, incorporation into proteins of; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Translation, genetic

(incorporation of unnatural amino acids in; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Protein engineering

(of aminoacyl-tRNA synthetases; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Glycosylation

(of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Genetic engineering

Molecular cloning

(synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Glycoproteins

RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)

(synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT tRNA

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(tyrosine-specific, variants, for introduction of amino acid analogs; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT Methanococcus jannaschii

(tyrosyl tRNA synthetase of; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 60063-85-6, Galacturonyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(Galacturonyltransferase, for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 9054-44-8, N-Acetylgalactosaminyltransferase
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
 (Uses)

(acetylgalactosaminyltransferase, for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)
4302-79-2 684302-80-5 684302-81-6 684302-82-7 684302-83-8

IT 684302-79-2 684302-80-5 684302-81-6 684302-82-7 684302-83-8 684302-84-9

RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); PRP (Properties); BIOL'(Biological study); USES (Uses) (amino acid sequence; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 61652-90-2DP, protein conjugates 70858-45-6DP, protein conjugates 81034-76-6DP, protein conjugates 84808-02-6DP, protein conjugates

RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)

(enzymic preparation in protein glycosidation; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 9023-45-4D, Tyrosyl tRNA synthetase, variants

RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological study); USES (Uses)

(for charging tRNA with amino acid analogs; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 57534-80-2, Glucuronic acid transferase 192588-73-1, Glycoprotein glucuronyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 56626-18-7, Fucosyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(fucosyltransferase, for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 9031-68-9, Galactosyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(galactosyltransferase, for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 9031-48-5, Glucosyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(glucosyltransferase, for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 7512-17-6, N-Acetylglucosamine

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(glycosidation of proteins at; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

TT 528-04-1 2956-16-3, UDP-galactose 3123-67-9,
 GDP-mannose 9054-94-8, β-1,4-Galactosyltransferase 83744-93-8
 135622-87-6 193099-05-7 202420-38-0 334993-76-9
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(glycosidation of proteins using; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

17041-36-0 21008-33-3 67315-18-8 685088-58-8

17041-36-0 21008-33-3 67315-18-8 685088-58-8
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(incorporation into proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 22888-49-9P

ΙT

RL: BUU (Biological use, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses)

(introduction into proteins of; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 9055-06-5, Mannosyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(mannosyltransferase, for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 684302-85-0 684302-86-1 684302-87-2 684302-88-3
RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological study); USES (Uses) (nucleotide sequence; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 66640-86-6, Biotin hydrazide 109653-47-6, Fluorescein hydrazide

RL: RCT (Reactant); RACT (Reactant or reagent)
 (protein modification with; synthetic glycosylation of proteins by
 incorporation of unnatural amino acids with novel reactive groups into
 protein)

IT 78-67-1, AIBN 122-00-9 128-08-5, N-Bromosuccinimide 1068-90-2, Diethyl acetamidomalonate 204716-07-4

RL: RCT (Reactant); RACT (Reactant or reagent)

(reactions of; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 321976-25-4, Sialyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(sialyltransferase, for glycosidation of proteins; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 56093-23-3,  $\alpha$ -1,2 Fucosyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

 $(\alpha-1,2$ -fucosyltransferase, glycosidation of proteins using; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 68247.-53-0,  $\alpha(1,3)$ -Fucosyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

 $(\alpha-1,3$ -fucosyltransferase, glycosidation of proteins using; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 111310-37-3, α(1,4)-Fucosyltransferase

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

 $(\alpha-1,4-fucosyltransferase, glycosidation of proteins using;$ 

synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

IT 7512-17-6, N-Acetylglucosamine

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(glycosidation of proteins at; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

RN 7512-17-6 HCAPLUS

CN D-Glucose, 2-(acetylamino)-2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT 528-04-1 2956-16-3, UDP-galactose

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(glycosidation of proteins using; synthetic glycosylation of proteins by incorporation of unnatural amino acids with novel reactive groups into protein)

RN 528-04-1 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate), P'-[2-(acetylamino)-2-deoxy- $\alpha$ -D-glucopyranosyl] ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 2956-16-3 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate), P'- $\alpha$ -D-galactopyranosyl ester (9CI) (CA INDEX NAME)

IT 109653-47-6, Fluorescein hydrazide

RL: RCT (Reactant); RACT (Reactant or reagent)
(protein modification with; synthetic glycosylation of proteins by
incorporation of unnatural amino acids with novel reactive groups into
protein)

RN 109653-47-6 HCAPLUS

CN Benzoic acid, 2-(3,6-dihydroxy-9H-xanthen-9-yl)-, hydrazide (9CI) (CA INDEX NAME)

L59 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2003:652522 HCAPLUS

DN 139:285735

ED Entered STN: 22 Aug 2003

TI Identification of Active-Site Inhibitors of MurG Using a Generalizable, High-Throughput Glycosyltransferase Screen

AU Helm, Jeremiah S.; Hu, Yanan; Chen, Lan; Gross, Ben; Walker, Suzanne

CS Department of Chemistry, Princeton University, Princeton, NJ, 08544, USA

SO Journal of the American Chemical Society (2003), 125(37), 11168-11169 CODEN: JACSAT; ISSN: 0002-7863

PB American Chemical Society

DT Journal

LA English

CC 1-5 (Pharmacology)

Section cross-reference(s): 7, 10

AB MurG is a glycosyltransferase involved in the biosynthesis of bacterial peptidoglycan. It is a potentially important antibiotic target, but no inhibitors of the enzyme have been reported. In general, inhibitors of glycosyltransferases have been difficult to design. Furthermore, no glycosyltransferase inhibitors have been identified through

high-throughput screening, perhaps because appropriate screens for glycosyltransferase inhibition have not been developed. In this manuscript, the authors describe the development of a high-throughput screen for MurG that was used to screen a 50 000 compound library for inhibitors. The screen, which can be generalized to other glycosyltransferases, led to the identification of a family of active-site directed MurG inhibitors. The family of inhibitors contains a five-membered heterocyclic core that appears to function as a diphosphate mimic with respect to the presentation of substituents. The authors discuss the implications of this result and the utility of the screen for identifying inhibitors of other glycosyltransferases.

ST MurG glycosyltransferase inhibitor identification high throughput screen

IT Antibacterial agents

Combinatorial library

Drug screening

High throughput screening

(identification of active-site inhibitors of MurG using a generalizable high-throughput glycosyltransferase screen in relation to antibacterial activity)

IT Enzyme functional sites

(inhibitor-binding; identification of active-site inhibitors of MurG using a generalizable high-throughput glycosyltransferase screen in relation to antibacterial activity)

IT Enzyme kinetics

(of inhibition; identification of active-site inhibitors of MurG using a generalizable high-throughput glycosyltransferase screen in relation to antibacterial activity)

IT 608143-47-1

RL: BUU (Biological use, unclassified); PAC (Pharmacological activity); BIOL (Biological study); USES (Uses)

(displacement ligand; identification of active-site inhibitors of MurG using a generalizable high-throughput glycosyltransferase screen in relation to antibacterial activity)

IT 60976-26-3, MurG glycosyltransferase

RL: BSU (Biological study, unclassified); BUU (Biological use,

unclassified); BIOL (Biological study); USES (Uses)

(identification of active-site inhibitors of MurG using a generalizable high-throughput glycosyltransferase screen in relation to antibacterial activity)

IT 312501-65-8

RL: PAC (Pharmacological activity); BIOL (Biological study) (identification of active-site inhibitors of MurG using a generalizable high-throughput glycosyltransferase screen in relation to antibacterial activity)

RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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- (26) Wong, K; Adv Exp Med Biol 1998, V456, P197 HCAPLUS
- (27) Zhang, Z; Annu Rev Pharmacol Toxicol 2002, V42, P209 HCAPLUS

### IT 608143-47-1

RL: BUU (Biological use, unclassified); PAC (Pharmacological activity); BIOL (Biological study); USES (Uses)

(displacement ligand; identification of active-site inhibitors of MurG using a generalizable high-throughput glycosyltransferase screen in relation to antibacterial activity)

RN 608143-47-1 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate), P'-[2-deoxy-2-[[[[(3',6'-dihydroxy-3-oxospiro[isobenzofuran-1(3H),9'-[9H]xanthen]-5yl)carbonyl]amino]acetyl]amino]-α-D-glucopyranosyl] ester (9CI) (CA
INDEX NAME)

- L59 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN
- AN 1996:367026 HCAPLUS
- DN 125:50470
- ED Entered STN: 25 Jun 1996
- TI Molecular cloning of the Golgi apparatus uridine diphosphate-N-acetylglucosamine transporter from Kluyveromyces lactis
- AU Abeijon, Claudia; Robbins, Phillips W.; Hirschberg, Carlos B.
- CS Dep. Biochem. Mol. Biol., Univ. Massachusetts Med. Cent., Worcester, MA, 01655, USA

Proceedings of the National Academy of Sciences of the United States of SO America (1996), 93(12), 5963-5968 CODEN: PNASA6; ISSN: 0027-8424 PB National Academy of Sciences DT Journal LΑ English CC 3-3 (Biochemical Genetics) Section cross-reference(s): 6, 10 AR The mannan chains of Kluyveromyces lactis mannoproteins are similar to those of Saccharomyces cerevisiae except that they lack mannose phosphate and have terminal  $\alpha 1\rightarrow 2$ -linked N- acetylglucosamine. The biosynthesis of these chains probably occurs in the human of the Golgi apparatus, by analogy to S. cerevisiae. The sugar donors, GDP-mannose and UDP-GlcNAc, must first be transported from the cytosol, their site of synthesis, via specific Golgi membrane transporters into the lumen where they are substrates in the biosynthesis of these mannoproteins. A mutant of K. lactis, mnn2-2, that lacks terminal Nacetylglucosamine in its mannan chains in vivo, has recently been characterized and shown to have a specific defect in transport of UDP-GlcNAc into the lumen of Golgi vesicles in vitro. We have now cloned the gene encoding the K. lactis Golgi membrane UDP-GlcNAc transporter by complementation of the mnn2-2 mutation. The mnn2-2 mutant was transformed with a genomic library from wild-type K. lactis in a pKD1-derived vector; transformants were isolated and phenotypic correction was monitored following cell surface labeling with fluorescein isothiocyanate conjugated to Griffonia simplicifolia II lectin, which binds terminal N-acetylglucosamine and a fluorescent activated cell sorter. A 2.4-kb DNA fragment was found to restore the wild-type lectin binding phenotype. Upon loss of the plasmid containing this fragment, reversion to the mutant phenotype occurred. The above fragment contained an open reading frame for a multitransmembrane spanning protein of 328 amino acids. contains a leucine zipper motif and has high homol. to predicted proteins from S. cerevisiae and C. elegans. In an assay in vitro, Golgi vesicles isolated from the transformant had reqained their ability to transport UDP-GlcNAc. Taken together, the above results strongly suggest that the cloned gene encodes the golgi UDP-GlcNAc transporter of K. lactis. ST Kluyveromyces Golgi uridine diphosphate acetylglucosamine transporter; uridine diphosphate acetylglucosamine transporter gene sequence TΤ Cell membrane (2.4-kb DNA fragment which was found to restore the wild-type lectin binding phenotype to Kluyveromyces lactis mnn2-2 mutant contained an open reading frame for a multitransmembrane spanning protein of 328 amino acids) IT Complementation, genetic (gene encoding the Kluyveromyces lactis Golgi membrane UDP-GlcNAc transporter has been cloned by complementation of the mnn2-2 mutation) IT Mutation (mnn2-2; gene encoding the Kluyveromyces lactis Golgi membrane UDP-GlcNAc transporter has been cloned by complementation of the mnn2-2 mutation) IT Golgi apparatus Kluyveromyces lactis Molecular cloning (mol. cloning of the Golgi apparatus uridine diphosphate

-N-acetylglucosamine transporter from Kluyveromyces lactis)

IT

Protein sequences

```
(of the Golgi apparatus uridine diphosphate-N-
        acetylglucosamine transporter from Kluyveromyces lactis)
IT
     Deoxyribonucleic acid sequences
        (of the Golgi apparatus uridine diphosphate-N-
        acetylglucosamine transporter gene from Kluyveromyces lactis)
ΙT
     Plasmid and Episome
        (upon loss of the plasmid containing the 2.4-kb DNA fragment which was
        found to restore the wild-type lectin binding phenotype to
        Kluyveromyces lactis mnn2-2 mutant, reversion to the mutant phenotype
        occurred)
IT
     Proteins, specific or class
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (UDP-N-acetylglucosamine-transporting, mol. cloning
        of the Golgi apparatus uridine diphosphate-N-
        acetylglucosamine transporter from Kluyveromyces lactis)
IT
     Conformation and Conformers
        (leucine zipper, protein which was found to restore the wild-type
        lectin binding phenotype to Kluyveromyces lactis mnn2-2 mutant contains
        a leucine zipper motif and has high homol. to predicted proteins from
        S. cerevisiae and C. elegans)
     178235-60-4
TT
     RL: PRP (Properties)
        (amino acid sequence; mol. cloning of the Golgi apparatus uridine
        diphosphate-N-acetylglucosamine transporter from
        Kluyveromyces lactis)
     528-04-1, Uridine diphosphate-N-
IT
     acetylglucosamine
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PRP (Properties); BIOL (Biological study)
        (mol. cloning of the Golqi apparatus uridine diphosphate
        -N-acetylglucosamine transporter from Kluyveromyces lactis)
IT
     177526-16-8, GenBank U48413
     RL: PRP (Properties)
        (nucleotide sequence; mol. cloning of the Golgi apparatus uridine
        diphosphate-N-acetylglucosamine transporter from
        Kluyveromyces lactis)
IT
     528-04-1, Uridine diphosphate-N-
     acetylglucosamine
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PRP (Properties); BIOL (Biological study)
        (mol. cloning of the Golqi apparatus uridine diphosphate
        -N-acetylglucosamine transporter from Kluyveromyces lactis)
RN
     528-04-1 HCAPLUS
CN
    Uridine 5'-(trihydrogen diphosphate), P'-[2-(acetylamino)-2-deoxy-\alpha-
    D-glucopyranosyl] ester (9CI) (CA INDEX NAME)
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ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN
L59
     1988:525962 HCAPLUS
AN
DN
     109:125962
ED
     Entered STN: 14 Oct 1988
     Metabolism of D-[U-14C] glucosamine in seedlings of Calluna vulgaris (L.)
TT
     Hull
     Piro, G.; Perotto, S.; Bonfante-Fasolo, P.; Dalessandro, G.
ΑU
     Dip. Biol., Univ. Stud. Lecce, Lecce, I-73100, Italy
CS
     Journal of Plant Physiology (1988), 132(6), 695-701
SO
     CODEN: JPPHEY; ISSN: 0176-1617
DT
     Journal
LA
    English
     11-2 (Plant Biochemistry)
CC
     Exogenous D-[U-14C] glucosamine was taken up by roots of C.
AB
     vulgaris seedlings, translocated throughout the plant and metabolized to
     N-acetyl-D-glucosamine, N-acetyl-D-
     glucosamine 6-phosphate, N-acetyl-D-glucosamine
     1-phosphate, UDP-N-acetyl-D-glucosamine and
     UDP-N-acetyl-D-galactosamine. The N-acetyl
     -D-hexosamine nucleosides acted as glycosyl donors of polymers containing N-
     acetyl-D-glucosamine and N-acetyl
     -D-galactosamine residues. The presence of N-acetyl-D-
     glucosamine residues at the surface of hair roots was shown by
     using wheat germ agglutinin linked to fluorescein isothiocyanate
     or colloidal gold as specific probes.
ST
     Calluna glucosamine metab
IT
    Root
        (acetylglucosamine residues at surface of, localization of)
IT
     Glycoproteins, biological studies
     RL: BIOL (Biological study)
        (glucosamine incorporation in, in Calluna vulgaris seedlings)
IT
     Calluna vulgaris
        (glucosamine metabolism in seedlings of)
IT
     Translocation
        (of glucosamine in Calluna vulgaris seedlings)
IT
     528-04-1, UDP-N-acetyl-D-glucosamine
     1746-32-3, N-Acetyl-D-glucosamine 6-phosphate
     6866-69-9 7277-98-7, UDP-N-acetyl
     -D-galactosamine 7512-17-6, N-Acetyl-D-
     glucosamine
    RL: FORM (Formation, nonpreparative)
        (formation of, from glucosamine, in Calluna vulgaris
        seedings)
IT
     3416-24-8, D-Glucosamine
```

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (metabolism of, in Calluna vulgaris seedlings) IT 528-04-1, UDP-N-acetyl-D-glucosamine 1746-32-3, N-Acetyl-D-glucosamine 6-phosphate 6866-69-9 7277-98-7, UDP-N-acetyl -D-galactosamine 7512-17-6, N-Acetyl-Dglucosamine RL: FORM (Formation, nonpreparative) (formation of, from glucosamine, in Calluna vulgaris seedings) 528-04-1 HCAPLUS RNUridine 5'-(trihydrogen diphosphate), P'-[2-(acetylamino)-2-deoxy- $\alpha$ -CN D-glucopyranosyl] ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 1746-32-3 HCAPLUS
CN D-Glucose, 2-(acetylamino)-2-deoxy-, 6-(dihydrogen phosphate) (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 6866-69-9 HCAPLUS CN D-Glucopyranose, 2-(acetylamino)-2-deoxy-, 1-(dihydrogen phosphate) (9CI) (CA INDEX NAME)

RN 7277-98-7 HCAPLUS

CN Uridine 5'-(trihydrogen diphosphate), P'-[2-(acetylamino)-2-deoxy- $\alpha$ -D-galactopyranosyl] ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 7512-17-6 HCAPLUS

CN D-Glucose, 2-(acetylamino)-2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

=> d his

L1

(FILE 'HOME' ENTERED AT 15:03:15 ON 18 OCT 2005)
SET COST OFF

FILE 'HCAPLUS' ENTERED AT 15:03:41 ON 18 OCT 2005

1 S US20050142629/PN OR US2003-784335#/AP, PRN

E KAHNE D/AU

L2 117 S E3-E7

e kahne s/au

L3 8 S E4, E5

E WALKER S/AU

L4 210 S E3

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L5
             20 S E17
               E WALKER SUE/AU
L6
             48 S E21
                SEL RN L1
    FILE 'REGISTRY' ENTERED AT 15:05:45 ON 18 OCT 2005
L7
            48 S E1-E48
L8
             4 S L7 AND (OC5-C6-C6 OR OC4-OC5-C6-C6-C6)/ES
L9
             2 S L8 AND OC5/ES
             1 S L9 AND NCNC3/ES
L10
L11
         75563 S (OC5-C6-C6 OR OC4-OC5-C6-C6-C6)/ES
L12
         1637 S L11 AND OC5/ES
L13
           103 S L12 AND (OC4 AND NCNC3)/ES
L14
            91 S L13 AND P/ELS
L15
            28 S L14 AND 8/NR
            17 S L15 NOT S/ELS
L16
             1 S L16 AND C38H38N4O23P2
L17
             2 S L8 AND OC4-OC5-C6-C6-C6/ES NOT L9
L18
             1 S L18 NOT N/ELS
L19
L20
             1 S 58-98-0
               E N-ACT5EYLGLUCOSAMINE/CN
               E N-ACETYLGLUCOSAMINE/CN
              1 S E3
L21
               E C38H38N4O23P2/MF
     FILE 'HCAOLD' ENTERED AT 15:16:10 ON 18 OCT 2005
L22
             0 S L17
     FILE 'HCAPLUS' ENTERED AT 15:16:14 ON 18 OCT 2005
L23
             2 S L17
L24
              0 S L19 AND L20 AND L21
L25
          6310 S L19
L26
         25115 S FLUORESCEIN?
L27
         25589 S L25,L26
L28
         2084 S L20
         16687 S UDP OR URID? (L) DIHYDROGEN (L) ?PHOSPH?
L30
        16657 S UDP OR URID? (L)?PHOSPH? (L)GLUCORON?
L31
          4962 S URID? (L) DIPHOSPH?
            40 S L27 AND L28-L31
L33
          5969 S L21
L34
          25484 S ?ACETYLGLUCOSAMIN? OR ?ACETY?(L)?GLUCOSAMIN?
L35
              9 S L32 AND L33, L34
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L36
             1 S 528-04-1
    FILE 'HCAPLUS' ENTERED AT 15:25:47 ON 18 OCT 2005
          1276 S L36
             7 S L37 AND L27
             1 S L38 NOT L35
               SEL DN AN L35 2 4 8 9
             4 S E1-E12 AND L35
             4 S L37 AND L40
             5 S L23, L41
            2 S L1-L6 AND L42
             5 S L42, L43
           2 S L44 AND GLCNAC
L45
L46
             5 S L44,L45
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FILE 'USPATFULL, USPAT2' ENTERED AT 15:29:27 ON 18 OCT 2005

L47 1 S L17

FILE 'USPATFULL, USPAT2' ENTERED AT 15:29:46 ON 18 OCT 2005

FILE 'HCAPLUS' ENTERED AT 15:29:53 ON 18 OCT 2005 SEL RN L46

	FILE	'REGIS	T	۱ Y۶	ENTE	RED	ΑT	15:3	0:24	ON	18	OCT	2005
L48		103	S	E13	-E119	5							
L49		4	S	L48	AND	OC4	1-00	25 - C6	-C6 <b>-</b>	C6/I	ES		
L50		1	s	L48	AND	L36	5						
L51		4	S	L48	AND	L10	),L	L7,L1	9,L2	0,L2	21		
L52		7	S	L49	-L51								
L53		6	S	L48	AND	(NC	CNC	AND	OC4	)/ES	S NO	OT L	52
L54		3	S	L53	NOT	(TF	[MY	DINE	OR	C6/I	ES)		
L55		10	S	L52	,L54								
L56		90	S	L48	TOM	L49	9-L5	55					
L57		3	S	L56	AND	(CE	3H16	NO9P	OR	C20I	1161	1204)	ı
L58		. 13	S	L55	,L57								

FILE 'HCAPLUS' ENTERED AT 15:36:01 ON 18 OCT 2005 L59 5 S L58 AND L46

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